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     3
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         APR 04
                STN AnaVist $500 visualization usage credit offered
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        MAY 10
                CA/CAplus enhanced with 1900-1906 U.S. patent records
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        MAY 11
                KOREAPAT updates resume
NEWS
     7
        MAY 19
                Derwent World Patents Index to be reloaded and enhanced
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        MAY 30
                IPC 8 Rolled-up Core codes added to CA/CAplus and
                USPATFULL/USPAT2
NEWS 9 MAY 30
                The F-Term thesaurus is now available in CA/CAplus
        JUN 02
NEWS 10
                The first reclassification of IPC codes now complete in
                 INPADOC
NEWS 11
         JUN 26
                TULSA/TULSA2 reloaded and enhanced with new search and
                and display fields
         JUN 28
NEWS 12
                Price changes in full-text patent databases EPFULL and PCTFULL
NEWS 13
         JUL 11 CHEMSAFE reloaded and enhanced
NEWS 14
         JUl 14
                FSTA enhanced with Japanese patents
NEWS 15
         JUl 19
                Coverage of Research Disclosure reinstated in DWPI
NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT
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MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.

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FILE 'MEDLINE' ENTERED AT 10:04:54 ON 05 AUG 2006

FILE 'AGRICOLA' ENTERED AT 10:04:54 ON 05 AUG 2006

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=> dup rem 11

PROCESSING COMPLETED FOR L1

27 DUP REM L1 (34 DUPLICATES REMOVED)
ANSWERS '1-5' FROM FILE MEDLINE
ANSWERS '6-7' FROM FILE BIOSIS
ANSWERS '8-27' FROM FILE CAPLUS

=> d his

L2

(FILE 'HOME' ENTERED AT 10:04:27 ON 05 AUG 2006)

FILE 'MEDLINE, AGRICOLA, DRUGU, JICST-EPLUS, CABA, BIOTECHNO, BIOSIS, CAPLUS, LIFESCI, BIOTECHDS, EMBASE, BIOENG, SCISEARCH' ENTERED AT 10:04:54 ON 05 AUG 2006

L1 61 S HOMOSERINE TRANSSUCCINYLASE

L2 27 DUP REM L1 (34 DUPLICATES REMOVED)

=> d 12 ibib abs total

L2 ANSWER 1 OF 27 MEDLINE on STN ACCESSION NUMBER: 2005203044 MEDLINE

DUPLICATE 3

DOCUMENT NUMBER: PubMed ID: 15838036

TITLE: Polyphosphate kinase protects Salmonella enterica from weak

organic acid stress.

Price-Carter Marian; Fazzio Thomas G; Vallbona Ester AUTHOR:

Ibanez; Roth John R

CORPORATE SOURCE: Department of Biology, University of Utah, Salt Lake City,

Utah 84112, USA.

CONTRACT NUMBER: GM34804 (NIGMS)

Journal of bacteriology, (2005 May) Vol. 187, No. 9, pp. SOURCE:

3088-99.

Journal code: 2985120R. ISSN: 0021-9193.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

200506 ENTRY MONTH:

Entered STN: 20 Apr 2005 ENTRY DATE:

> Last Updated on STN: 22 Jun 2005 Entered Medline: 21 Jun 2005

AB Mutants of Salmonella enterica lacking polyphosphate kinase (ppk) grow poorly in the presence of the weak organic acids acetate, propionate, and This sensitivity is corrected by methionine and seems to result from destabilization of MetA (homoserine

transsuccinylase), the first enzyme in methionine biosynthesis. The MetA protein is known to be sensitive to thermal inactivation, and ppk mutants are more sensitive to heat-induced methionine auxotrophy. Peroxide increases the sensitivity of ppk mutants to both heat and acid

and may oxidatively damage (carbonylate) destabilized MetA. While acid appears to impair methionine biosynthesis, it leads to derepression of MetA and may inhibit growth by causing toxic accumulation of denatured protein. This is supported by the observation that the overexpression of MetA in ppk mutants causes acid sensitivity that is not corrected by methionine. We propose that polyphosphate acts as a chemical chaperone that helps refold MetA and/or may stimulate proteolysis of toxic denatured protein. The instability of MetA protein may provide a metabolic fuse that blocks growth under conditions that denature proteins; the sensitivity of this fuse is modulated by polyphosphate.

DUPLICATE 7 ANSWER 2 OF 27 MEDLINE on STN

ACCESSION NUMBER: 2000391212 MEDLINE DOCUMENT NUMBER: PubMed ID: 10913262

TITLE: Enzyme-catalyzed acylation of homoserine: mechanistic

characterization of the Haemophilus influenzae met2-encoded

homoserine transacetylase.

Born T L; Franklin M; Blanchard J S AUTHOR:

CORPORATE SOURCE: Department of Biochemistry, Albert Einstein College of

Medicine, Bronx, NY 10461, USA.

CONTRACT NUMBER: AI33696 (NIAID)

GM19514 (NIGMS)

Biochemistry, (2000 Jul 25) Vol. 39, No. 29, pp. 8556-64. Journal code: 0370623. ISSN: 0006-2960. SOURCE:

United States PUB. COUNTRY:

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200008

Entered STN: 24 Aug 2000 ENTRY DATE:

> Last Updated on STN: 24 Aug 2000 Entered Medline: 15 Aug 2000

AB The first unique step in bacterial and plant methionine biosynthesis involves the acylation of the gamma-hydroxyl of homoserine. Haemophilus influenzae, acylation is accomplished via an acetyl-CoA-dependent acetylation catalyzed by homoserine transacetylase.

The activity of this enzyme regulates flux of homoserine into multiple biosynthetic pathways and, therefore, represents a critical control point for cell growth and viability. We have cloned homoserine transacetylase from H. influenzae and present the first detailed enzymatic study of this enzyme. Steady-state kinetic experiments demonstrate that the enzyme utilizes a ping-pong kinetic mechanism in which the acetyl group of acetyl-CoA is initially transferred to an enzyme nucleophile before subsequent transfer to homoserine to form the final product, O-acetylhomoserine. The maximal velocity and V/K(homoserine) were independent of pH over the range of values tested, while V/K(acetyl)(-)(CoA) was dependent upon the ionization state of a single group exhibiting a pK value of 8.6, which was required to be protonated. Solvent kinetic isotope effect studies yielded inverse effects of 0.75 on V and 0.74 on V/K(CoA) on the reverse reaction and effects of 1.2 on V and 1.7 on V/K(homoserine) on the forward reaction. Direct evidence for the formation of an acetyl-enzyme intermediate was obtained using rapid-quench labeling studies. On the basis of these observations, we propose a chemical mechanism for this important member of the acyltransferase family and contrast its mechanism with that of homoserine transsuccinylase.

L2 ANSWER 3 OF 27 MEDLINE on STN DUPLICATE 8

ACCESSION NUMBER: 2000042598 MEDLINE DOCUMENT NUMBER: PubMed ID: 10572016

TITLE: Enzyme-catalyzed acylation of homoserine: mechanistic

characterization of the Escherichia coli metA-encoded

homoserine transsuccinylase.

AUTHOR: Born T L; Blanchard J S

CORPORATE SOURCE: Department of Biochemistry, Albert Einstein College of

Medicine, Bronx, New York 10461, USA.

CONTRACT NUMBER: AI33696 (NIAID)

GM19514 (NIGMS)

SOURCE: Biochemistry, (1999 Oct 26) Vol. 38, No. 43, pp. 14416-23.

Journal code: 0370623. ISSN: 0006-2960.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199912

ENTRY DATE: Entered STN: 13 Jan 2000

Last Updated on STN: 13 Jan 2000 Entered Medline: 17 Dec 1999

AB The first unique step in bacterial and plant methionine biosynthesis involves the activation of the gamma-hydroxyl of homoserine. In Escherichia coli, this activation is accomplished via a succinylation reaction catalyzed by homoserine transsuccinylase. The activity of this enzyme is closely regulated in vivo and therefore represents a critical control point for cell growth and viability. We have cloned homoserine transsuccinylase from E. coli and present the first detailed enzymatic study of this enzyme. Steady-state kinetic experiments demonstrate that the enzyme utilizes a ping-pong kinetic mechanism in which the succinyl group of succinyl-CoA is initially transferred to an enzyme nucleophile before subsequent transfer to homoserine to form the final product, O-succinylhomoserine. The maximal velocity, V/K(succinyl)(-)(CoA), and V/K(homoserine) all exhibited a bell-shaped pH dependence with apparent pK's of 6.6 and approximately 7.9. The enzyme was inhibited by iodoacetamide in a pH-dependent manner, with an apparent pK of the group being inactivated of 6.4. This suggests the presence of an active site cysteine which forms a succinyl-cysteine intermediate during enzymatic turnover. Solvent kinetic isotope effect studies yielded inverse effects of 0.7 on V and 0.61 on V/K in the reverse reaction only. On the basis of these observations, we propose a detailed chemical mechanism for this important member of the acyltransferase

L2 ANSWER 4 OF 27 MEDLINE on STN DUPLICATE 9

ACCESSION NUMBER: 95173116 MEDLINE DOCUMENT NUMBER: PubMed ID: 7868613

TITLE: Heat shock-dependent transcriptional activation of the metA

gene of Escherichia coli.

AUTHOR: Biran D; Brot N; Weissbach H; Ron E Z

CORPORATE SOURCE: Department of Molecular Microbiology and Biotechnology,

Tel-Aviv University, Israel.

SOURCE: Journal of bacteriology, (1995 Mar) Vol. 177, No. 5, pp.

1374-9.

Journal code: 2985120R. ISSN: 0021-9193.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199503

ENTRY DATE: Entered STN: 7 Apr 1995

Last Updated on STN: 6 Feb 1998 Entered Medline: 27 Mar 1995

In Escherichia coli, the growth rate at elevated temperatures is AΒ controlled by the availability of endogenous methionine, which is limited because of the temperature sensitivity of the metA gene product, homoserine transsuccinylase (HTS). In order to determine the relationship between this control mechanism and the heat shock response, we estimated the cellular levels of HTS during heat shock by Western (immunoblot) analysis and found an increase following induction by temperature shift and by addition of ethanol or cadmium ions. The elevated level of HTS was a result of transcriptional activation of the This activation was heat shock dependent, as it did not take metA gene. place in rpoH mutants, and probably specific to the metA gene, as another gene of the methionine regulon (metE) was not activated. These results suggest a metabolic link between the two systems that control the response of E. coli to elevated temperatures: the metA gene, which codes for the enzyme responsible for regulating cell growth as a function of temperature elevation (HTS), is transcriptionally activated by the heat shock response.

L2 ANSWER 5 OF 27 MEDLINE on STN DUPLICATE 12

ACCESSION NUMBER: 76024802 MEDLINE DOCUMENT NUMBER: PubMed ID: 1100601

TITLE: Growth rate of Enterobacteriaceae at elevated temperatures:

limitation by methionine.

AUTHOR: Ron E Z

SOURCE: Journal of bacteriology, (1975 Oct) Vol. 124, No. 1, pp.

243-6.

Journal code: 2985120R. ISSN: 0021-9193.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 197512

ENTRY DATE: Entered STN: 13 Mar 1990

Last Updated on STN: 13 Mar 1990 Entered Medline: 29 Dec 1975

AB The effect of elevated temperatures on growth rate was studied in five strains of Enterobacteriaceae. In all the strains tested a shift to the elevated temperature resulted in an immediate decrease in growth rate which was due to limitation in the availability of endogenous methionine. The first biosynthetic enzyme of the methionine pathway-homoserine transsuccinylase-was studied in extracts of Aerobacter aerogenes, Salmonella typhimurium, and Escherichia coli and was shown to be

temperature sensitive in all of them.

L2 ANSWER 6 OF 27 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

ACCESSION NUMBER: 1983:73430 BIOSIS

DOCUMENT NUMBER: PREV198324073430; BR24:73430

TITLE: USE OF A RECOMBINANT PLASMID CONTAINING THE MET-A GENE OF

ESCHERICHIA-COLI TO STUDY GROWTH OF ENTEROBACTERIACEAE AT

ELEVATED TEMPERATURES.

AUTHOR(S): MICHAELI S [Reprint author]; RON E Z

CORPORATE SOURCE: TEL-AVIV UNIV

SOURCE: (1982) pp. P95. INTERNATIONAL UNION OF MICROBIOLOGICAL

SOCIETIES. 13TH INTERNATIONAL CONGRESS OF MICROBIOLOGY; BOSTON, MASS., USA, AUG. 8-13, 1982. XIV+182P. AMERICAN SOCIETY FOR MICROBIOLOGY: WASHINGTON, D.C., USA. PAPER.

ISBN: 0-914826-44-1.

DOCUMENT TYPE: Book

Conference; (Meeting)

FILE SEGMENT: BR
LANGUAGE: ENGLISH

L2 ANSWER 7 OF 27 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

ACCESSION NUMBER: 1982:171714 BIOSIS

DOCUMENT NUMBER: PREV198273031698; BA73:31698

TITLE: CONSTRUCTION AND PHYSICAL MAPPING OF PLASMIDS CONTAINING

THE META GENE OF ESCHERICHIA-COLI K-12.

AUTHOR(S): MICHAELI S [Reprint author]; RON E Z; COHEN G

CORPORATE SOURCE: DEP MICROBIOL, GEORGE S WISE FACULTY LIFE SCI, TEL-AVIV

UNIV, TEL-AVIV, ISRAEL

SOURCE: Molecular and General Genetics, (1981) Vol. 182, No. 2, pp.

349-354.

CODEN: MGGEAE. ISSN: 0026-8925.

DOCUMENT TYPE: Article

FILE SEGMENT: BA
LANGUAGE: ENGLISH

AB Plasmids containing the metA gene (which codes for homoserine transsuccinylase of the methionine biosynthetic pathway) of E. coli K-12 were constructed in vitro using plasmid pBR322 as the cloning vehicle and λmetA transducing phage as the source of metA DNA. EcoRI digests of pBR322 and λmetA20 were jointed by ligase and plasmids carrying the metA gene were selected after transformation in a metA deletion strain. Recombinant DNA molecules contained 1 pBR322 fragment and 1 λmetA20 fragment of 12.2 kb [kilobases] which was present in either of 2 possible orientations. Plasmids constructed by BamH1 digestion of λmetA2 contained a single bacterial DNA fragment of 5.8 kb inserted in the tet gene. Insertion of the metA fragment led to loss of resistance to tetracycline in one orientation and partial

resistance in the opposite orientation.

L2 ANSWER 8 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2005:1239041 CAPLUS

DOCUMENT NUMBER: 144:2275

TITLE: Construction of microorganism containing recombinant

homoserine transsuccinylase with

altered feedback sensitivity and recombinant

S-adenosylmethionine synthetase with reduced activity

for the production of methionine

INVENTOR(S): Bestel-Corre, Gwenaeelle Anne Lise; Chateau, Michel;

Figge, Rainer Martin; Raynaud, Celine; Soucaille,

Philippe Noeel Paul

PATENT ASSIGNEE(S): Metabolic Explorer, Fr.

SOURCE: PCT Int. Appl., 72 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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                                              WO 2005-EP52180
     WO 2005108561
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PRIORITY APPLN. INFO.:
                                               WO 2004-IB1901
                                                                  A 20040512
                           CASREACT 144:2275; MARPAT 144:2275
OTHER SOURCE(S):
     The present invention relates to the use of recombinant homoserine
     transsuccinylase with altered sensitivity to feedback inhibitors
     S-adenosylmethionine and methionine (MetA*) and optionally, recombinant
     S-adenosylmethionine synthetase with reduced activity (MetK*) for the
     production of methionine, its precursors or derivs. thereof. More
     specifically, the authors isolated Escherichia coli mutants containing
     homoserine transsuccinylase which show decreased
     feedback-sensitivity towards S-adenosylmethionine and methionine. E. coli
     mutants containing S-adenosylmethionine synthetase with reduced activity were
     also isolated. Construction of E. coli strains for the production of
     O-succinylhomoserine or methionine by combining feed-back resistant MetA
     alleles with MetK alleles with decreased activity is described. Fermentation
of
     E. coli production strains and anal. of yield is reported.
                                 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
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                                  RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 9 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
                           2005:1220814 CAPLUS
ACCESSION NUMBER:
                           143:474228
DOCUMENT NUMBER:
TITLE:
                           Construction of microbial recombinant
                           homoserine transsuccinylase with
                           altered feedback sensitivity and S-adenosyl methionine
                           synthetase with reduced activity for the production of
                           methionine
INVENTOR(S):
                           Bestel-Corre, Gwenaeelle; Chateau, Michel; Figge,
```

Rainer Martin; Raynaud, Celine; Soucaille, Philippe

PATENT ASSIGNEE(S): Metabolic Explorer, Fr. SOURCE: PCT Int. Appl., 46 pp.

Noel Paul

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

of

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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    WO 2005108561
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             SN, TD, TG
PRIORITY APPLN. INFO.:
                                             WO 2004-IB1901
                                                                 A 20040512
OTHER SOURCE(S):
                         CASREACT 143:474228; MARPAT 143:474228
     The present invention relates to the use of recombinant homoserine
     transsuccinylase with altered feedback sensitivity (MetA*) and
     eventually, recombinant S-adenosyl methionine synthetase with reduced
     activity (MetK*) for the production of methionine, its precursors or derivs.
     thereof. More specifically, Escherichia coli mutants containing
    homoserine transsuccinylase with decreased feedback
     sensitivity towards methionine and S-adenosylmethionine were isolated. E.
     coli mutants containing S-adenosyl methionine synthetase with reduced activity
     were also isolated. Construction of E. coli strains for the production of
     O-succinylhomoserine or methionine by combined feed-back resistant MetA
     alleles with MetK alleles with decreased activity is described. Fermentation
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E. coli production strains and anal. of yield is reported.

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ANSWER 10 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 4
ACCESSION NUMBER:
                        2004:371078 CAPLUS
DOCUMENT NUMBER:
                        140:387796
                        Methionine and SAM feedback-resistant
TITLE:
                        homoserine transsuccinylases with
                        modified C-terminus
                        Leonhartsberger, Susanne; Pfeiffer, Kerstin;
INVENTOR(S):
                         Winterhalter, Christoph; Bauer, Brigitte
```

PATENT ASSIGNEE(S): Consortium fuer Elektrochemische Industrie G.m.b.H., Germany

SOURCE: PCT Int. Appl., 33 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

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		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
			ΙE,	SI,	FI,	RO,	CY,	TR,	BG,	CZ,	EE,	HU,	SK					
	CN	1705	751			Α		2005	1207		CN 2	2003-	8010	1894		2	0031	016
	JP	2006	5035	68		Т2		2006	0202	1	JP 2	2004-	5458	67		2	0031	016
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AB The invention relates to a homoserine transsuccinylase , which exhibits reduced sensitivity towards L-methionine or SAM in comparison with a homoserine transsuccinylase wild-type enzyme, whereby the latter comprises an amino acid sequence containing a TyrGlnXaaThrPro sub-sequence, the Thr of said sub-sequence lying between positions 285 and 310 of the amino acid sequence. The inventive homoserine transsuccinylase is characterized in that in comparison with the wild-type enzyme at least 2 amino acids are modified, said modification taking place in the Thr of the sub-sequence or in the C-terminal. Thus, exts. of E. coli containing metA gene mutants were analyzed for homoserine transsuccinylase activity in the presence of 1 mM Met or SAM. The wild-type enzyme retains 2% and 0.5% activity, resp. One of the mutants exhibited 95% activity under these circumstances. The Ki for Met was 16 mM and for SAM 9 mM.

L2 ANSWER 11 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 5

ACCESSION NUMBER:

2004:349595 CAPLUS

DOCUMENT NUMBER:

140:370810

TITLE:

Feedback-resistant homoserine

transsuccinylase mutants, microorganisms

producing them, and their use in production of

methionine and SAM

INVENTOR(S):

Winterhalter, Christoph; Leonhartsberger, Susanne;

Pfeiffer, Kerstin; Bauer, Brigitte

PATENT ASSIGNEE(S):

Consortium fuer Elektrochemische Industrie G.m.b.H.,

Germany

SOURCE:

Ger. Offen., 22 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.		KIND	DATE	APPLICATION NO.	DATE			
		-						
DE 1024743	7	A1	20040429	DE 2002-10247437	20021011			
WO 2004035	517	A2	20040429	WO 2003-EP10978	20031002			
WO 2004035	517	A3	20040617					
W: CA	CN, J	P, RU, US	;					
RW: AT	BE, B	G, CH, CY	CZ, DE,	DK, EE, ES, FI, FR, GB,	GR, HU, IE,			
IT	LU, M	C, NL, PI	RO, SE,	SI, SK, TR				
EP 1549754		A2	20050706	EP 2003-767502	20031002			
R: AT	BE, C	I, DE, DE	K, ES, FR,	GB, GR, IT, LI, LU, NL,	SE, MC, PT,			
IE	SI, F	I, RO, CY	TR, BG,	CZ, EE, HU, SK				

20051130 CN 2003-80101208 20031002 20060622 JP 2004-544072 20031002 DE 2002-10247437 A 20021011 WO 2003-EP10978 W 20031002 CN 1703517 · A JP 2006516092 T2 PRIORITY APPLN. INFO.:

Homoserine transsuccinylase, which contains at least AB one mutation compared to a homoserine transsuccinylase wild type enzyme and compared to the wild type enzyme shows a reduced sensitivity to L-methionine or SAM is disclosed. The wild-type enzyme contains a partial sequence AspGlyXaaXaaXhrGlyAlaPro between residues 90 and 115 and a partial sequence TyrGlnXaaThrPro between residues 285 and The mutations comprise an amino acid exchange of Asp in AspGlyXaaXaaXaaThrGlyAlaPro or an amino acid exchange of Tyr in TyrGlnXaaThrPro. Thus, the Y294C mutant of E. coli MetA exhibits 96% activity in the presence of 1 mM Met while the wild-type enzyme is almost totally inhibited. The Ki for Met in the mutant is 11 mM, for Met in the wild-type, 0.05 mM. The same mutant show 92% activity in the presence of 1 mM SAM and a Ki of 10 mM, while the wild-type enzyme shows negligible activity and Ki of 0.2 mM.

ANSWER 12 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 6

ACCESSION NUMBER: 2000:342198 CAPLUS

DOCUMENT NUMBER: 133:3756

L-methionine and its preparation with transgenic TITLE:

Escherichia coli mutants with defective repressor and

enhanced homoserine transsuccinylase

activity

Usuta, Yoshihiro; Kurahashi, Osamu INVENTOR(S):

PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan Jpn. Kokai Tokkyo Koho, 23 pp. SOURCE:

CODEN: JKXXAF

DOCUMENT TYPE: Patent Japanese LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

KIND DATE APPLICATION NO. DATE PATENT NO. JP 2000139471 A2 20000523 JP 1998-326717 19981117 RITY APPLN. INFO.: JP 1998-326717 19981117 PRIORITY APPLN. INFO.: Described is a method of manufacturing L-methionine by cultivating a Escherichia

coli mutant with defective repressors (gene metJ), enhanced homoserine transsuccinylase (gene metA) activity, and, optionally, decreased S-adenosyl methionine synthetase activity. Furthermore, the mutants may also have the enhanced activities of cystathionine- γ -synthase and aspartokinase-homoserine dehydrogenase II. Also claimed are the S-adenosyl methionine synthetase (metK) mutants with substitution mutations at 27-Arg→Cys, 296-Ile→Ser, 298-Pro→Leu, or a combination of them. The mutants are free of the synergistic inhibition by L-methionine and S-adenosyl methionine. Production of L-methionine with improved efficiency by using the Escherichia coli mutants was demonstrated.

ANSWER 13 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 10

ACCESSION NUMBER: 1989:568215 CAPLUS

111:168215 DOCUMENT NUMBER:

Nucleotide sequence of the metA gene encoding TITLE:

homoserine trans-succinylase in Escherichia coli

Duclos, B.; Cortay, J. C.; Bleicher, F.; Ron, E. Z.; AUTHOR(S):

Richaud, C.; Saint Girons, I.; Cozzone, A. J.

CORPORATE SOURCE: Lab. Biol. Mol., Univ. Lyon, Villeurbanne, 69622, Fr.

Nucleic Acids Research (1989), 17(7), 2856 SOURCE:

CODEN: NARHAD; ISSN: 0305-1048

DOCUMENT TYPE: Journal LANGUAGE: English

AB The nucleotide sequence of the 927 bp long metA gene encoding homoserine transsuccinylase is presented. The deduced amino acid sequence indicates a protein of 35,673 daltons in good agreement with the predicted mol. mass of the polypeptide. The last 103 codons were part of an unidentified open reading frame immediately upstream of the aceB gene.

L2 ANSWER 14 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 11

ACCESSION NUMBER: 1984:524135 CAPLUS

DOCUMENT NUMBER: 101:124135

TITLE: Expression of the metA gene of Escherichia coli K-12

in recombinant plasmids

AUTHOR(S): Michaeli, Shulamit; Ron, Eliora Z.

CORPORATE SOURCE: George S. Wise Fac. Life Sci., Tel Aviv Univ., Tel

Aviv-Jaffa, Israel

SOURCE: FEMS Microbiology Letters (1984), 23(2-3), 125-9

CODEN: FMLED7; ISSN: 0378-1097

DOCUMENT TYPE: Journal LANGUAGE: English

AB The expression of the metA gene for homoserine transsuccinylase [9030-70-0] was studied in wild-type and in deregulated strains of E. coli K-12 carrying the gene on multicopy plasmids. The mol. weight of the product synthesized by the metA gene was 40,000; the whole enzyme consisted of 2 subunits. In deregulated strains (i.e., those carrying a metJ mutation), the activity of the metA gene was increased 2-fold. Thus, even when metA is cloned onto a multicopy plasmid, it is under the neg. control of the regulatory metJ gene.

L2 ANSWER 15 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:513390 CAPLUS

DOCUMENT NUMBER: 141:66285

TITLE: Virulence genes and proteins from Pseudomonas

aeruginosa and Klebsiella pneumoniae, and their

therapeutic, diagnostic and vaccine use

INVENTOR(S): Cosson, Pierre; Kohler, Thilo; Benghezal, Mohammed;

Marchetti, Anna; Van Delden, Christian

PATENT ASSIGNEE(S): University of Geneva, Switz.

SOURCE: U.S. Pat. Appl. Publ., 43 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

	PAT	ENT 1	NO.			KIN	D	DATE		1	APPL	ICAT:	ION	NO.		D	ATE		
US 2004122212				A1 20040624			US 2002-324967						20021220						
	US 6974680			B2 20051213															
	CA	2510	474			AA 20040708			CA 2003-2510474						20031219				
	WO 2004057018				A2 20040708			WO 2003-CH836						20031219					
	WO 2004057018				A3 20040902														
		W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,	
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			ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	
			TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG

20040714 AU 2003-286074 AU 2003286074 A1 20031219 20050928 EP 2003-776748 A2 EP 1578788 20031219 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK US 2005-536606 US 2006171959 20060803 A1 A 20021220 US 2002-324967 PRIORITY APPLN. INFO.: WO 2003-CH836 W 20031219

AB The present invention is based on the discovery of 46 genes, VIR1-VIR46, when mutated lower the virulence of a gram-neg. bacterium, and can be used in new antimicrobial therapeutic strategies. Particularly, 19 mutants from Pseudomonas (MUT1-MUT19) and 27 from Klebsiella (MUT20-MUT46) were analyzed in Dictyostelium discoideum host system, and shown to encode products that are implicated in virulence. The identification of these genes therefore allows attenuated microorganisms to be produced. Furthermore, the genes or their encoded products can be used to identify antimicrobial drugs, diagnostic methods for the identification of a pathogen-associated disease, and in the manufacture of vaccines.

REFERENCE COUNT: 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 16 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:1008670 CAPLUS

DOCUMENT NUMBER: 142:88744

TITLE: Probing the active site of homoserine

trans-succinylase

AUTHOR(S): Rosen, Ran; Becher, Doerte; Buettner, Knut; Biran,

Dvora; Hecker, Michael; Ron, Eliora Z.

CORPORATE SOURCE: Department of Molecular Microbiology and

Biotechnology, Faculty of Life Sciences, Tel-Aviv

University, Tel-Aviv, 69978, Israel FEBS Letters (2004), 577(3), 386-392

CODEN: FEBLAL; ISSN: 0014-5793

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

SOURCE:

AB Homoserine trans-succinylase is the first enzyme in methionine biosynthesis of Escherichia coli and catalyzes the activation of homoserine via a succinylation reaction. The in vivo activity of this enzyme is subject to tight regulation by several mechanisms, including repression and activation of gene expression, feedback inhibition, temperature regulation and proteolysis. This complex regulation reflects the key role of this enzyme in bacterial metabolism Here, the authors demonstrate—using proteomics and high-resolution mass spectrometry—that succinyl is covalently bound to one of the two adjacent lysine residues at positions 45 and 46. Replacing these lysine residues by alanine abolished the enzymic activity. These findings position the lysine residues, one of which is conserved, at the active site.

REFERENCE COUNT: 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 17 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1985:40804 CAPLUS

DOCUMENT NUMBER: 102:40804

TITLE: Regulatory region of the metA gene of Escherichia coli

K-12

AUTHOR(S): Michaeli, Shulamit; Mevarech, Moshe; Ron, Eliora Z. CORPORATE SOURCE: George S. Wise Fac. Life Sci., Tel Aviv Univ., Tel

Aviv-Jaffa, 69978, Israel

SOURCE: Journal of Bacteriology (1984), 160(3), 1158-62

CODEN: JOBAAY; ISSN: 0021-9193

DOCUMENT TYPE: Journal LANGUAGE: English

AB Transcription of the metA gene of E. coli K-12 is from a promoter which is

under methionine control and is located next to a region which has an extensive sequence homol. with the operator regions of the metBL and metF genes. However, in the metA gene, there is a 2nd transcription start point which is located 74 base pairs upstream and which is independent of the intracellular methionine concentration

ANSWER 18 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN L2

1983:84243 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 98:84243

Level of polyamines in Escherichia coli carrying the TITLE:

metA gene on a multicopy plasmid

AUTHOR(S): Michaeli, Shulamit; Rozenhak, Sonia; Ron, Eliora Z.

CORPORATE SOURCE: Dep. Microbiol., Tel-Aviv Univ., Tel Aviv-Jaffa,

Israel

SOURCE: Advances in Polyamine Research (1983), 4, 519-20

CODEN: APYRD9; ISSN: 0160-2179

DOCUMENT TYPE: Journal English LANGUAGE:

Strains of E. coli with elevated level of intracellular methionine were obtained by the introduction. of multicopy plasmids containing the metA gene,

which codes for homoserine transsuccinylase

[9030-70-0], the 1st enzyme in the methionine [63-68-3] pathway. One of the plasmids obtained which contained the metA gene was pMA-3. Strains carrying this plasmid were overproducers of methionine. In the presence of elevated intracellular methionine concns., there was an increase in spermidine [124-20-9] content that was concomitant with a decrease in the level of putrescine [110-60-1]; this resulted in a significant change in the ratio of spermidine-to-putrescine.

ANSWER 19 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN L2

1982:469190 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 97:69190

Mechanisms involved in the increased sensitivity of TITLE:

Escherichia coli to microcin 15m at 42°C

AUTHOR(S): Aguilar, Alfredo; Perez-Diaz, Jose C.; Asensio, Carlos

Inst. Enzimol. Patol. Mol., Madrid, Spain CORPORATE SOURCE: Current Microbiology (1982), 7(2), 83-6 SOURCE:

CODEN: CUMIDD; ISSN: 0343-8651

DOCUMENT TYPE: Journal LANGUAGE: English

E. coli Cells show a markedly increased sensitivity to the antibiotic microcin 15m when briefly treated at 42° as compared to the effect at 37°. Furthermore, mutants resistant to the microcin at 37° become sensitive at 42° at microcin concns. that are inactive at 37°. This effect can be overcome by L-methionine. The mechanism involved seems to be based on an apparent inactivation of the homoserine-O-transsuccinylase activity. As previously established, this enzyme suffers a reversible partial inactivation when the cells are shifted to 42° and the action of the microcin at this temperature seems to bring this process to a virtually irreversible stage. In mixed cultures of the microcin-producing strain and 1 E. coli stain sensitive to the antibiotic, a much stronger growth inhibition of the latter strain was observed at 42° than at 37° .

ANSWER 20 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1977:548496 CAPLUS

87:148496 DOCUMENT NUMBER:

TITLE: Norleucine accumulation by a norleucine-resistant

mutant of Serratia marcescens

AUTHOR(S): Kisumi, Masahiko; Sugiura, Masaki; Chibata, Ichiro CORPORATE SOURCE: Res. Lab. Appl. Biochem., Tanabe Seiyaku Co., Ltd.,

Osaka, Japan

SOURCE: Applied and Environmental Microbiology (1977), 34(2), 135 - 8

CODEN: AEMIDF; ISSN: 0099-2240

DOCUMENT TYPE:

LANGUAGE: English

A norleucine-resistant mutant was derived from an isoleucine-valine auxotroph of a leucine accumulator of S. marcescens. norleucine-resistant mutant could accumulate norleucine from norvaline in the medium without the addition of methionine, which antagonized norleucine. This mutant constitutively formed homoserine-O-transsuccinylase.

ANSWER 21 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1975:454709 CAPLUS

DOCUMENT NUMBER:

83:54709

TITLE:

SOURCE:

Methionine biosynthesis in isolated Pisum sativum

mitochondria

AUTHOR(S): CORPORATE SOURCE: Clandinin, Michael T.; Cossins, Edwin A. Dep. Bot., Univ. Alberta, Edmonton, AB, Can. Phytochemistry (Elsevier) (1974), 13(3), 585-91 CODEN: PYTCAS; ISSN: 0031-9422

DOCUMENT TYPE:

Journal

LANGUAGE: English

Homocysteine-dependent transmethylases utilizing 5methyltetrahydropteroylglutamic acid and S-adenosylmethionine as Me donors were examined in (NH4)2SO4 fractions prepared from isolated mitochondria of pea cotyledons. Substantial levels of 5-methyltetrahydropteroylglutamate transmethylase (I) [9033-23-2] were detected, the catalytic properties of this enzyme were similar to those of a previously reported enzyme present in cotyledon exts. The mitochondrial I had an apparent Km of 25 μM for the Me donor, was saturated with homocysteine at 1 mM and was inhibited 50% by L-methionine at 2.5 mM. At similar concns. of Me donor the mitochondrial S-adenosylmethionine methyltransferase was not saturated Mitochondrial prepns. were capable of synthesizing substantial amts. of S-adenosylmethionine but lacked ability to form S-methylmethionine. Significant levels of β -cystathionase, cystathionine- γ synthase, L-homoserine transacetylase, and L-homoserine transsuccinylase were detected in the isolated mitochondria. activity of the enzymes of homocysteine biosynthesis was not affected by L-methionine in vitro. Thus, pea mitochondria are able to catalyze the synthesis of methionine [63-68-3] de novo.

ANSWER 22 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1974:45506 CAPLUS

DOCUMENT NUMBER:

80:45506

TITLE:

Control of homoserine-O-transsuccinylase in a

methionine-requiring mutant of the blue-green alga

Anacystis nidulans

AUTHOR(S):

Delaney, S. F.; Dickson, A.; Carr, N. G.

CORPORATE SOURCE:

Dep. Biochem., Univ. Liverpool, Liverpool, UK Journal of General Microbiology (1973), 79(Pt. 1),

SOURCE:

89-94

CODEN: JGMIAN; ISSN: 0022-1287

DOCUMENT TYPE:

Journal

LANGUAGE: English

The regulation of the 1st step in methionine biosynthesis, homoserine-O-transsuccinylase, has been examined in a methionine-requiring mutant of A. nidulans. No evidence of derepression of the biosynthesis of this enzyme was found even under conditions of acute methionine starvation. End product inhibition of the enzyme by homoserine, cystathionine, and methionine was demonstrated, and shown, in the latter case, to be of an allosteric nature. The lack of transcription control of this enzyme is discussed as an example of a general phenomenon in this group of microorganisms.

ANSWER 23 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1972:56522 CAPLUS

DOCUMENT NUMBER:

76:56522

TITLE:

Regulation of the methionine feedback-sensitive enzyme

in mutants of Salmonella typhimurium

AUTHOR(S):

SOURCE:

Lawrence, David A.

CORPORATE SOURCE:

Lab. Enzymol., CNRS, Gif-sur-Yvette, Fr. Journal of Bacteriology (1972), 109(1), 8-11

CODEN: JOBAAY; ISSN: 0021-9193

DOCUMENT TYPE:

Journal English

LANGUAGE:

Assay of the first enzyme unique to methionine biosynthesis, homoserine-O-transsuccinylase, in metJ and metK regulatory mutants of S. typhimurium showed that synthesis of the enzyme was derepressed seven- and four-fold, resp. The possibility of noncoordinate regulation of the methionine enzymes is discussed. In metA feedback resistant mutants the enzyme activity can be inhibited in vitro by 10 mM S-adenosylmethionine but not by 10 mM L-methionine. The synergistic inhibition found for the wild-type enzyme is not effective in these latter mutants.

ANSWER 24 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1972:55595 CAPLUS

DOCUMENT NUMBER:

76:55595

TITLE:

Miscellaneous procedures involved in transsulfuration

AUTHOR(S):

Flavin, Martin

CORPORATE SOURCE:

Lab. Biochem., Natl. Heart Lung Inst., Bethesda, MD,

USA

SOURCE:

Methods Enzymol. (1971), Volume 17, Issue Pt. B,

450-3. Editor(s): Colowick, S. P. Academic: New

York, N. Y. CODEN: 18HWA8

DOCUMENT TYPE:

Conference English

LANGUAGE:

Cystathionine β -synthase, assay of mixts. of β - and γ-cystathionases, β-cystathionase and cystathionine

γ-synthase from Neurospora, and homoserine transsuccinylase are reviewed. 17 refs.

ANSWER 25 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1971:484622 CAPLUS

DOCUMENT NUMBER:

75:84622

TITLE:

Growth rate of Escherichia coli at elevated

temperatures. Reversible inhibition of homoserine

trans-succinylase

AUTHOR(S):

Ron, Eliora Z.; Shani, M.

CORPORATE SOURCE:

Dep. Microbiol., Tel Aviv Univ., Tel Aviv, Israel

SOURCE:

Journal of Bacteriology (1971), 107(2), 397-400

CODEN: JOBAAY; ISSN: 0021-9193

DOCUMENT TYPE:

Journal

LANGUAGE: English A shift from 37 to 44° decreased the activity of crude or partially

purified E. coli homoserine transsuccinylase (I) prepns. This effect was rapid and immediately reversible. The change in I activity induced by elevated temperature may involve breaking of H or

hydrophobic bonds, since urea at 37° caused a similar reversible

lowering of I activity.

ANSWER 26 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

1970:528144 CAPLUS

DOCUMENT NUMBER:

73:128144

TITLE:

The mechanism of ethionine toxicity to Escherichia

AUTHOR(S):

Miller, Kathryn L.; Martin, William Randolph

CORPORATE SOURCE: Dep. of Microbiol., Univ. of Chicago, Chicago, IL, USA

SOURCE: Proceedings of the Society for Experimental Biology

and Medicine (1970), 135(2), 311-16

CODEN: PSEBAA; ISSN: 0037-9727

DOCUMENT TYPE: Journal LANGUAGE: English

AB Toxic effects of ethionine on E. coli in a glucose-salts medium are manifested by growth inhibition and loss of viability. Valine-14C incorporation and β -galactosidase studies indicate that ethionine rapidly inhibits protein synthesis. The methionine precursors, homocysteine and cystathionine, as well as methionine itself, reverse all ethionine effects while O-succinylhomoserine and homoserine do not. Homoserine O-transsuccinylase, the first enzyme unique for methionine biosynthesis, is inhibited by ethionine while the analog does not inhibit cystathionine synthetase or methionine transfer RNA synthetase. Data indicate that the initial effect of ethionine on E. coli is to act as an inhibitor of homoserine O-transuccinylase resulting in a severe depletion of methionine available for protein and other cellular synthesis.

L2 ANSWER 27 OF 27 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1971:431670 CAPLUS

DOCUMENT NUMBER: 75:31670

TITLE: Genetical study of the feedback-sensitive enzyme of

methionine synthesis in Salmonella typhimurium

AUTHOR(S): Chater, K. F.; Rowbury, R. J.

CORPORATE SOURCE: Genet. Dep., Univ. Birmingham, Birmingham, UK

SOURCE: Journal of General Microbiology (1970), 63(Pt. 1),

111-20

CODEN: JGMIAN; ISSN: 0022-1287

DOCUMENT TYPE: Journal LANGUAGE: English

AB The homoserine-O-transsuccinylase activity of 3 methionine excreting mutants of S. typhimurium was examined In 1 the enzyme was resistant to inhibition by methionine or its analog α -methylmethionine, while in the other 2 feedback inhibition was normal. The failure of methionine in the 1st was attributed to failure to penetrate the cells or to an alteration of homoserine-O-transsuccinylase such that it was not sensitive to feedback inhibition.

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